Prevalence of Medication Errors and the Associated Factors: A Prospective Observational Study Among Cancer Patients at Mbarara Regional Referral Hospital

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Background: Medication error is one of the most common medical errors in the practice of modern medicine. Among cancer patients receiving chemotherapy, medication errors can be potentially harmful given the narrow therapeutic index, complex dosing, and toxic nature of anti-cancer drugs.

Objective: This study aimed to determine the incidence and factors associated with medication errors among cancer patients.

Methods: The study was a prospective observational study carried out at the cancer unit of Mbarara Regional Referral Hospital, Southwestern Uganda. The study included 110 participants, both adults and children receiving chemotherapy. The study was carried out for a period of five months from January to May 2020. A checklist was used to collect patient, medication, and disease information to identify the prescription, transcription, dispensing, and administration errors.

Results: Of the 110 participants, 52 (47.3%) experienced a total of 78 medication errors (MEs). Of these, 33 (42.31%) were prescription errors, 29 (37.18%) administration errors, 9 (11.54%) transcription errors, and 7 (8.97%) dispensing errors. In the adjusted logistic regression of factors associated with medication errors, urban residents (aOR, 4.59; 95% CI, 1.08, 19.53, $p=0.039$) and educated participants (at secondary level) (aOR, 10.51; 95% CI, 1.43, 77.14, $p=0.021$) had a significantly higher risk of experiencing medication errors. Participants treated with alkylating agents (aOR, 2.87; 95% CI, 1.07, 7.72, $p=0.036$) had a greater risk of experiencing medication errors when compared to other classes of chemotherapy.

Conclusion: The incidence of medication errors among cancer patients was high in Mbarara Regional Referral Hospital. Prescription errors were the most common type of error followed by administration errors, and dispensing errors were the least common. Residence, education level, and alkylating agent chemotherapy were significantly associated with occurrence of medication errors.

Keywords: prevalence, medication error, associated factors, cancer, Mbarara

Introduction

Medication error is one of the most common medical errors in the practice of modern medicine.¹ According to © 2021 National Coordinating Council for Medication Error Reporting and Prevention, All Rights Reserved, medication errors are defined as:

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any preventable event that may cause or lead to inappropriate medication use or patient harm while the medication is in the control of the health care professional, patient, or consumer. Such events may be related to professional practice, health care products, procedures, and systems.¹

There is an increasing incidence of both adult and pediatric cancer cases in developing countries, thus, there is equally a rise in the use of anti-cancer drugs as one of the cancer treatments.² However, with the increase in drug use, there is also an increasing chance of medication errors which can occur at any stage of prescription, transcription, dispensing, and administration of anti-cancer drugs resulting in disability, prolonged hospitalization, and mortality.³ Chemotherapy-related medication errors can be potentially harmful given the narrow therapeutic index, complex dosing, and toxic nature of anti-cancer drugs especially among the already immune-compromised cancer patients.¹ A review of data about medication errors among cancer patients indicated that 1–3% of patients experienced a medication error during treatment.⁴ In the USA, the incidence of medication errors among cancer patients has been reported at 7.1% among adults and 18.8% among pediatric patients.⁵

Concisely, medication errors impose great economic impact on the healthcare system, society, and patients. The impact on health organizations, health workers, and patients include prolonged hospitalization, loss of trust, and loss of productivity respectively.⁶

Data reviewed about medication errors in Africa showed that 13% to 76% of prescriptions for general patients had a medication error.⁷ A study in Nigeria revealed that prescription medication error was the most common medical error at 95.2% among medical personnel.⁸ In Uganda, 18% of health care professionals disclosed having made medication errors while treating patients.⁹ Medication errors are therefore a common occurrence among inpatient and outpatient encounters. There has been no similar research conducted in the study area on incidence of medication errors and specifically data about cancer-related medication errors. Accordingly, this study aimed to determine the incidence and factors associated with medication errors in a cancer unit in Uganda.

Methods

Study Design and Setting

The study was conducted at the cancer unit of Mbarara Regional Referral Hospital, Southwestern Uganda. The

ward has a bed capacity of 20 beds for adults and 18 beds for pediatric cancer patients with two oncology specialists, one pharmacist, and seven nurses. The prospective observational study was carried out among patients on cycle one and two of treatment who were being treated at cancer unit of MRRH from January 2020 to May 2020.

Study Participants

All 110 patients who were receiving cycle one and two chemotherapy treatment at cancer unit of MRRH during the study period, who fulfilled the inclusion and exclusion criteria were studied.

Inclusion Criteria

The study included:

- Patients of all age groups.
- Patients with definitive diagnosis of cancer who were receiving chemotherapy or other adjuvant cancer drugs in cycle one and two of treatment.
- Patients who were able to provide a signed consent form (for adults) and consent obtained from caretakers of pediatric patients. Assent was also obtained from children and their care takers who were able to understand the study.

Exclusion Criteria

- Patients without a definitive cancer diagnosis and not receiving chemotherapy treatment did not participate in the study.
- Individuals that were unable to provide consent or assent were also excluded from the study.

Data Collection

A checklist was developed based on the American Society of Health-System Pharmacists¹⁰ classification of medication errors and used to document observed errors from each patient's medication while admitted to the cancer unit. The checklist included different drug prescription, transcription, dispensing, and administration errors.

Prescription errors were assessed by comparison of patient prescriptions with standard treatment protocol from Uganda Cancer Institute treatment guidelines and Physicians' Cancer Chemotherapy Drug Manual¹¹ with anti-cancer drugs that determined whether the prescriptions fulfilled the five-rights of appropriate drug use and correct frequency with duration of treatment regimen. The five-rights of drug use include right drug for the right

patient with the right indication at the right dose using the right route of administration. Transcription errors were determined from assessment of any differences in the patient prescription and drug transcription forms which are the basis for drug preparation.

Dispensing errors were identified as labeling errors, wrong drug, wrong doses, and wrong patients. Administration errors were differences in time of administration, frequency of administration, and route of administration. Following prescription for the study patient, the patient's prescription was assessed on whether the prescription followed the protocol for particular cancer treatment for the drug, calculated dose per body surface area, route, frequency, and duration prescribed.

After prescription the drugs are transcribed to the dispensing form that is used for preparation and dispensing of chemotherapy. The dispensing form was assessed for transcription error in comparison with the prescription. Following preparation, the drugs are labeled with patient name, drug name, dose as prescribed. Errors were identified based on the protocol and prescription. During drug administration after dispensing, the patient was observed for route, time, duration, and rate of administration. The errors were identified from the comparison with the treatment protocol.

To minimize the possible consequent bias, the errors identified prospectively were not corrected, except in few cases when investigators believed there was potentially life-threatening medication errors, when intervention was made during medical rounds without letting the health care workers know it was part of the study's finding.

Accuracy of the data collected was ensured through training of data collection assistants, close supervision of data collection process, and giving feedback in case of difficulties encountered. In addition, a pretest which involved five percent (9 patients) of the estimated sample size population was conducted and the feedback used to improve the data collection tools. The results obtained were not used in the final analysis of the study.

Data Processing and Analysis

The information obtained from individual patients' checklists was transferred to Excel. The collected data were analyzed using Stata version 12. Descriptive statistics were used to analyze frequency and percentage of medication errors among different variables. Chi-square d test was used to determine distribution and types of medication error among different categories of patients.

The association between each independent variable and medication error was analyzed using univariate logistic regression to obtain crude odds ratios. P-value less than 0.05 was considered significant. All variables with $P < 0.25$ were fit into a multivariate logistic regression model to control for confounders and obtain adjusted odds ratios.

Ethical Considerations

This study was conducted in accordance with the Declaration of Helsinki.¹² The study was approved by MUST Research and Ethics Committee (REC) with an approval letter Ref: MUREC 1/7-2020. Consent was sought before recruitment of adult patients while parental consent and child assent was obtained for children below eighteen years.

Results

Participant Characteristics

Of the 110 participants enrolled in this study, 67 (60.9%) were male, 34 (30.9%) were above 65 years of age, 68 (61.8%) were Munyankole ethnicity, 54 (49.1%) were rural residents, 80 (72.7%) participants had no comorbidities, 57 (51.8%) of the cancers diagnosed were not staged, 47 (42.7%) had not attained any formal education, and 66 (60.0%) were underweight. More than half of the participants were diagnosed with carcinomas, 67 (60.9%), while 63 (57.3%) were treated using alkylating agents as chemotherapy (Table 1).

Prevalence of Medication Errors

Overall, 52 (47.3%) participants had at least one medication error when all the four sub-categories of medication errors were considered (Figure 1).

Types of Medication Errors

The study showed that 33 (42.31%) prescription errors were the most commonly occurring, followed by 29 (37.18%) which were administration errors (Table 2). There were 9 (11.54%) transcription errors and 7 (8.97%) dispensing errors; the least common. Of the prescription errors, wrong dose prescribed (18) and wrong frequency (10) were the most common, while of the administration errors, wrong infusion rate (17) of administration was the most common.

Table 1 Participant Characteristics (n=110)

Characteristics	Level	Frequency (%)
Age category (years)	<18	28 (25.5)
	18–49	27 (24.5)
	50–64	21 (19.1)
	≥65	34 (30.9)
Sex	Male	67 (60.9)
	Female	43 (39.1)
Tribe	Munyankole	68 (61.8)
	Baganda	7 (6.4)
	Mukiga	21 (19.1)
	Others	14 (12.7)
Residence	Urban	18 (16.4)
	Peri-urban	38 (34.5)
	Rural	54 (49.1)
Level of education	None	47 (42.7)
	Primary	41 (37.3)
	Secondary	10 (9.1)
	Tertiary	12 (10.9)
Body mass index (BMI)	Underweight	66 (60.0)
	Normal weight	26 (23.6)
	Overweight	17 (15.5)
	Obese	1 (0.9)
Comorbidity	None	80 (72.7)
	DM	3 (2.7)
	Hypertension	8 (7.3)
	HIV	10 (9.1)
	Others	9 (8.2)
Stage of cancer	II	7 (6.4)
	III	15 (13.6)
	IV	31 (28.2)
	Not staged	57 (51.8)
Cancer diagnosis	Carcinoma	67 (60.9)
	Sarcoma	15 (13.6)
	Leukemia	9 (8.2)
	Lymphoma	8 (7.3)
	Others	10 (9.1)

(Continued)

Table 1 (Continued).

Characteristics	Level	Frequency (%)
Chemotherapy drugs	Alkylating agents	63 (57.3)
	Anti-metabolites	25 (22.7)
	Anthracyclines	53 (48.2)
	Topoisomerase inhibitors	2 (1.8)
	Plant alkaloid	46 (41.8)
	Corticosteroids	14 (12.8)
	Other treatments	50 (45.5)

Bivariate Logistic Regression for Factors Associated with Medication Error

In the unadjusted analysis, data showed that urban residents (OR, 3.69; 95% CI, 1.15–11.79, $p=0.028$) were more likely to experience medication errors compared to rural and peri-urban residents. Among participants who attained education, participants with secondary level education (OR, 5.97; 95% CI, 1.11–32.1, $p=0.037$) were more likely to experience medication errors. Participants receiving alkylating agents were also more likely to experience medication errors (OR 3.04; 95% CI, 1.37–6.70, $p=0.006$) (Table 3).

Multivariate Analysis of the Factors Associated with MEs

Multivariate analysis showed that urban residents were more likely to experience medication errors [adjusted

Prevalence of medication errors

Figure 1 Prevalence of medication errors among cancer patients at MRRH.

Table 2 Description of the Types and Categories of Medication Errors Observed Among Cancer Patients at Mbarara Regional Referral Hospital

Types of Medication Error (Frequency)	Description (Frequency)
Prescription errors (33)	Drug dose (18)
	Wrong frequency prescribed (10)
	Wrong route prescribed (2)
	Others (2)
	Wrong drug (1)
Administration error (29)	Wrong infusion rate (17)
	Others (12)
Transcription error (9)	Wrong dose (3)
	Wrong time (3)
	Others (3)
Dispensing error (7)	Wrong frequency (4)
	Wrong dose (3)

odds ratio (aOR), 4.59; 95% CI, 1.08, 19.53, $p=0.039$) compared to rural and peri-urban residents. Participants with secondary education were associated with more medication errors (aOR 10.51; 95% CI, 1.43, 77.14, $p=0.021$). Compared to participants receiving any other type of chemotherapy, participants on alkylating agents were more likely to experience a medication error (aOR 2.87; 95% CI, 1.07, 7.72, $p=0.036$) (Table 4).

Discussion

We conducted a study with the aim of documenting the prevalence of medication errors and the associated factors among cancer inpatients and outpatients at Mbarara Regional Referral Hospital from January to May 2020. Results revealed that during that period, 78 medication errors occurred, at a prevalence of 47.3% and most commonly, prescription errors (42.3%). The categories of wrong dose errors (54.5%) and frequency errors (30.4%) occurred more commonly. Administration errors were also frequent, unlike dispensing errors. Both bivariate and multivariate analyses showed that urban residents, participants who attained secondary level education, and those who received alkylating agents were more likely to experience medication errors.

Our results are in line with previous studies, for example; a study in South India documented a prevalence of 41.6%, with prescription errors as the most common type of error.¹³ The most common error was dose-related with 79 (22.83%) cases of overdosing. However, the current prevalence is much higher compared to the finding of a study in Brazil that reported a prevalence of 6.24%¹⁴ as well as another study in France that documented prevalence at 5.2%.¹⁵ These differences are probably because of the better health care systems and lower prescriber or nurse to patient ratio in middle income and high-income countries compared to our study that was conducted in a low-income country.

On the contrary, the current prevalence (47.3%) is lower than that reported in Pakistan (62%).¹⁶ This notable difference is probably due to the difference in methodology since the study included lack of effectiveness as one of the other errors involving drugs. The lower error prevalence observed in our study could also be due to human error since the prescription process is highly human dependent, characterized by hand written prescription and drug process orders without secondary verification procedures. These differences could also be attributed to the use of CPOE, prescription validation systems and training received by health care workers, a crucial component that is lacking in a developing country with poor health-care systems.

Our study also revealed that prescription errors were the most common error type, a finding similar to previous studies conducted in the United States,¹⁷ Spain,¹⁸ and Brazil.¹⁴ The high frequency of errors during the prescription process could be due to the shortage of oncology specialists to supervise the prescription process. The absence of updated specific cancer treatment protocols for individual patient files, accompanied by inadequate continuous training of prescribers on efficient use of protocols, could also have led to increased error occurrence during prescriptions. This is because treatment protocols guide prescribers on choice of drug regimen during prescriptions, hence errors could arise from their incorrect use and unavailability.¹⁹

Administration errors contributed 37.2% of all medication errors, compared to previous studies that documented 32% and 39% in the United Kingdom and Ethiopia respectively.^{20,21} However, a study from Iran indicated 26.5%, possibly due to the higher number of nursing staff involved in patient care compared to those in this study.²² Higher prevalence of administration errors was

Table 3 Bivariate Logistic Regression Analysis of the Independent Factors with MEs

Characteristics			Medication Errors			
	Level		No (n=58)	Yes (n=52)	Unadjusted OR (95% CI)	P-value
			No. (%)	No. (%)		
Age category (years)	<18		16 (27.6)	12 (23.1)	1	
	18–49		14 (24.1)	13 (25)	1.24 (0.43, 3.58)	0.694
	50–64		9 (15.5)	12 (23.1)	1.78 (0.57, 5.58)	0.324
	≥65		19 (32.76)	15 (28.9)	1.05 (0.35, 1.58)	0.921
Sex	Female		23 (41.4)	20 (36.5)	1	
	Male		35 (60.3)	32 (61.5)	1.05 (0.48, 2.26)	0.898
Residence	Rural		29 (50.0)	22 (42.3)	1	
	Peri-urban		24 (41.4)	16 (30.8)	0.88 (0.38, 2.04)	0.763
	Urban		5 (8.6)	14 (26.9)	3.69 (1.15, 11.79)	0.028
Level of education	None		29 (50.0)	17 (32.7)	1	
	Primary		22 (37.9)	21 (40.4)	1.62 (0.69, 3.8)	0.259
	Secondary		2 (3.5)	7 (13.5)	5.97 (1.11, 32.1)	0.037
	Tertiary		5 (8.6)	7 (13.5)	2.39 (0.65, 8.7)	0.187
Comorbidity	None		43 (74.1)	37 (71.2)	1	
	DM		1 (1.7)	1 (1.9)	1.16 (0.07, 19.23)	0.916
	Hypertension		6 (10.3)	3 (5.8)	0.58 (0.14, 2.49)	0.464
	HIV		5 (8.6)	5 (9.6)	1.16 (0.31, 4.32)	0.823
	Others		2 (3.4)	7 (13.5)	2.32 (0.54, 9.94)	0.256
Stage of cancer	II		5 (8.6)	2 (3.8)	1	
	III		9 (15.5)	6 (11.5)	2.5 (0.36, 17.5)	0.356
	IV		10 (17.2)	21 (40.4)	4.55 (0.75, 27.42)	0.099
	Not staged		34 (58.6)	23 (44.2)	1.64 (0.29, 9.19)	0.572
Cancer diagnosis	Carcinoma	No	25 (43.1)	18 (34.6)	1	
		Yes	33 (56.9)	34 (65.4)	1.43 (0.66, 3.10)	0.363
	Sarcoma	No	51 (87.9)	44 (84.6)	1	
		Yes	7 (12.1)	8 (15.4)	1.32 (0.44, 3.95)	0.614
	Leukemia	No	50 (86.2)	50 (96.2)	1	
		Yes	8 (13.8)	2 (3.9)	0.25 (0.05, 1.24)	0.089
	Lymphoma	No	55 (94.8)	47 (90.4)	1	
		Yes	3 (5.2)	5 (9.6)	1.95 (0.44, 8.6)	0.377
	Others	No	51 (87.9)	49 (94.2)	1	
		Yes	7 (12.1)	3 (5.8)	0.45 (0.11, 1.82)	0.261

(Continued)

Table 3 (Continued).

Characteristics			Medication Errors			
	Level		No (n=58)	Yes (n=52)	Unadjusted OR (95% CI)	P-value
			No. (%)	No. (%)		
Chemotherapy drugs	Alkylating agents	No	32 (55.2)	15 (28.9)	1	
		Yes	26 (44.8)	37 (71.2)	3.04 (1.37, 6.70)	0.006
	Anti-metabolites	No	44 (75.9)	41 (78.9)	1	
		Yes	14 (24.1)	11 (21.2)	0.84 (0.34, 2.07)	0.709
	Anthracyclines	No	28 (48.3)	29 (55.8)	1	
		Yes	30 (51.7)	23 (44.2)	0.74 (0.34, 1.6)	0.433
	Topoisomerase inhibitors	No	57 (98.3)	51 (98.1)	1	
		Yes	1 (1.7)	1 (1.92)	1.12 (0.07, 18.3)	0.938
	Plant alkaloid	No	34 (58.6)	30 (57.7)	1	
		Yes	24 (41.4)	22 (42.3)	1.03 (0.48, 2.2)	0.921
	Corticosteroids	No	51 (87.9)	45 (86.5)	1	
		Yes	7 (12.1)	7 (13.5)	1.13 (0.36, 3.5)	0.827
	Other treatments	No	29 (50)	31 (59.6)	1	
		Yes	29 (50.0)	21 (40.4)	0.68 (0.32, 1.4)	0.313

registered in a study conducted in United States (56%),⁵ probably due to the inclusion of preparation errors under the administration process which differs from our study. Rate of infusion errors (58.6%) were the most common in

this category under drug administration. This prevalence is higher than that reported by a study in Turkey which had an outcome of 39.7%² and 35.2% from Iran.²² This finding could be attributed to the use of infusion pumps that are

Table 4 Multivariate Logistic Regression Analysis of the Independent Factors with MEs

Characteristics	Level		Adjusted Odds Ratio	
			aOR (95% CI)	P-value
Residence	Rural		Ref	
	Peri-urban		0.89 (0.34, 2.32)	0.811
	Urban		4.59 (1.08, 19.53)	0.039
Level of education	None		Ref	
	Primary		1.54 (0.59, 4.02)	0.376
	Secondary		10.51 (1.43, 77.14)	0.021
	Tertiary		1.37 (0.24, 7.6)	0.719
Chemotherapy drugs	Alkylating agents	No	Ref	
		Yes	2.87 (1.07, 7.72)	0.036

Notes: 1) Adjusted odds ratios were expressed using exponentiated coefficients; 95% confidence intervals in brackets; 2) bolded figures indicate statistically significant associations.

Abbreviations: OR, unadjusted odds ratio; aOR, adjusted odds ratio.

necessary for standardizing drug delivery rates, which are absent in our study setting.

The other category of errors was under transcription with 11.54% contribution to the total errors. Wrong frequency transcribed, wrong doses, and other transcription errors contributed 33.3%. The total error is lower than the 21% reported by Ford et al and Schwappach and Wernli.^{23,24} The observed transcription errors can be attributed to multitasking required by the few pharmacy staff required to transcribe, prepare, and dispense chemotherapy. In addition, WHO links such errors to poor handwriting and lack of communication between health workers.²⁵

Dispensing errors (8.97%) were the least common as the drugs are mostly dispensed by trained pharmacists that frequently double-check before dispensing which reduces the risk of errors.¹⁶ The minor occurrences can be attributed to workload, and long work shifts since the pharmacist to patient ratio is very low.

Among the factors identified to be related to occurrence of medication error was urban residence, ie, urban residents experienced more medication errors compared to those in semi-urban and rural areas. This could be explained by an observation made that people living in urban areas preferred to be outpatients which compelled them to alter the drug infusion rate in order to leave hospital on the same day hence reducing the monitoring given to these patients compared to the inpatients.

In addition to residence, more errors were reported among the educated participants compared to the uneducated, with the greatest risk being among participants who had attained education at secondary school level. This is possibly linked to the settlement areas, ie, the educated are commonly found in urban areas. However, this is contrary to most studies which have revealed that more errors have been observed among the uneducated in studies carried out in community pharmacies due to inability to understand instructions and comprehend health education.²⁶

Participants who were treated with alkylating agents experienced more medication errors compared to other classes of chemotherapy treatment. This differs from the observation made in a study conducted among the pediatric population where anti metabolites were more associated with medication errors at 39.5% and alkylating agents at 14%.²⁷ However, WHO links errors associated with alkylating agents to the necessity to alter doses which is often not done as required.²⁸

The strengths of this study include prospective data collection which captured errors that could have been missed by retrospective method and using the patient study population which provided less biased information on occurrence of errors compared to health workers. However, some of the study limitations include a short study period of only five months, small pediatric study population, preparation process stage was not considered, and exclusion of documentation error under administration process.

Conclusion

Almost half of the patients at Mbarara Regional Referral Hospital experienced at least one medication error. The prevalence was comparable to other previous studies from developing countries. Prescription errors were the most common type of error which is comparable to other previous studies; mainly due to wrong dose being prescribed. Administration errors were second most frequent which was mostly due to wrong rate of infusion. Residence, education level, and alkylating agents used as chemotherapy were significantly associated with occurrence of medication errors.

Prescribers should take extra precautions when they prescribe chemotherapy by adhering to treatment protocols. Additionally, CPOE should be adopted to reduce human error related to hand-written prescriptions. Hospitals should avail chemotherapy infusion pumps to reduce errors resulting from rate of infusion. Patient health education is highly recommended in order to prevent medication errors caused by patients' lack of knowledge regarding complexity of their treatment.

Abbreviations

aOR, adjusted odds ratio; OR, odds ratio; DM, diabetes mellitus; HTN, hypertension; HIV, human immunodeficiency virus; CPOE, computerized physician order entry; MRRH, Mbarara University of Science and Technology; WHO, World Health Organization.

Data Sharing Statement

All the data supporting the conclusions of this article are included within this manuscript.

Ethics Approval and Consent to Participate

The study protocol was approved by the Research Ethics Committee (REC) at Mbarara University of Science and Technology as the Institutional review board for

authorization to conduct the research (MUREC 1/7-2020). Informed consent was sought before recruitment of adult patients while parental or guardian consent was obtained on behalf of children below 18 years. All methods used in the study were in accordance with regulations and guidelines of the Research Ethics Committee and cancer unit of Mbarara Regional Referral Hospital.

Consent for Publication

All authors agreed to submission of this manuscript for publication in addition to the consent to publish which was included in the informed consent form which attained ethical and participant approval.

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Author Contributions

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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Disclosure

The authors declare that they have no conflicts of interest for this work.

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